Genus or species specific detection of bacteria - by hybridising RNA with labelled specific probe after elongation in presence of specific or general primer

Patent Assignee: BOEHRINGER MANNHEIM GMBH; ROCHE DIAGNOSTICS GMBH Inventors: BERNER S; KESSLER C; KRUSE-MUELLER C; LUDWIG W; RUEGER R; RUGER R; SCHLEIFER

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Patent Family (11 patents, 17 countries)

Patent Number	Kind	Date	Application Number	Kind	Date	Update	Type
EP 480289	A	19920415	EP 1991116745	A	19911001	199216	В
DE 4038804	A	19920416	DE 4032024	A	19901009	199217	E
			DE 4038804	A	19901205		
CA 2052668	A	19920410	CA 2052668	A	19911002	199226	E
ZA 199108024	A	19920729	ZA 19918024	A	19911008	199235	E
JP 4258299	A	19920914	JP 1991257519	A	19911004	199243	E
JP 1994040839	B2	19940601	JP 1991257519	A	19911004	199420	E
EP 480289	B1	19970312	EP 1991116745	A	19911001	199715	E
DE 59108610	G	19970417	DE 59108610	A	19911001	199721	E
,			EP 1991116745	A	19911001		
ES 2101706	T3	19970716	EP 1991116745	A	19911001	199735	E
CA 2052668	C	20001219	CA 2052668	A	19911002	200103	E
US 6225094	B1	20010501	US 1991772026	A	19911008	200126	E
			US 199373985	A	19930608		

Priority Application Number (Number Kind Date): DE 4032024 A 19901009; DE 4038804 A 19901205

Patent Details

Patent Number	Kind	Language	Pages	Drawings	Filing Notes
EP 480289	A	DE	9	0	
Regional Designated States, Original	AT BE CH DE DK ES FR GB GR IT LI LU NL SE				
DE 4038804	A	DE	5		
CA 2052668	A	EN			
ZA 199108024	A	EN	47		
JP 4258299	A	JA	6		
JP 1994040839	B2	JA	6		Based on OPI patent JP 04258299
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Regional Designated States, Original	AT BE CH DE DK ES FR GB GR IT LI LU NL SE				
DE 59108610	G	DE			Application EP

			1991116745
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ES 2101706	Т3	ES	Application EP 1991116745
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CA 2052668	С	EN	
US 6225094	B1	EN	Continuation of application US 1991772026

Alerting Abstract: EP A

Genus and/or species specific detection of bacteria in test fluids comprises (1) hybridising bacterial RNA with a primer (I) which is (a) complementary to a genus/species specific region of RNA of certain bacteria, or to a highly conserved region of RNA of bacteria in general, but (b) at its 3'-end is not complementary to RNA of bacteria of other genera or species. (2) elongating the primer in presence of a polymerase and the 4 deoxyribonucleotides, opt with simultaneous or subsequent labelling of the elongation product (EP). (3) denaturing EP and hybridising with a genus/species specific oligonucleotide (II), then (4) detecting hybridisation from the label.

(I) is pref. complementary to RNA of one genus only and (II) will hybridise with RNA of only particular species within that genus. (I) and (II) are derived from r-, m- or t-RNA and can carry different labels.

Polymerisation is pref. with AMV or MMuLV reverse transcriptase, and before hybridisation with (II) EP are sepd. according to size by gel electrophoresis or ascending TLC. The label in EP may be one member of a specific binding pair, with the other partner being bound to a solid phase. Hybridisation with (II) is then done on this phase.

USE/ADVANTAGE - Method can distinguish between RNA differing in only a few bases. (I) and (II) provide two selection stages, so the danger of false assignments is reduced. Method is used e.g. to differentiate between harmful and harmless species

International Classification (Main): C12P-019/34, C12Q, C12Q-001/68 (Additional/Secondary): G01N, G01N-033/68

US Classification, Issued: 435091510, 435091200, 435091100, 435006000

Original Publication Data by Authority

Canada

Publication Number: CA 2052668 A (Update 199226 E)

Publication Date: 19920410

Assignee: BOEHRINGER MANNHEIM GMBH (BOEF)

Inventor: LUDWIG W SCHLEIFER K H KESSLER C RUEGER R STERN A

Language: EN

Application: CA 2052668 A 19911002 (Local application)
Priority: DE 4032024 A 19901009 DE 4038804 A 19901205

Original IPC: C12Q-1/68(A)

Current IPC: C12Q-1/68(A)|CA 2052668 C (Update 200103 E)

Publication Date: 20001219

Assignee: ROCHE DIAGNOSTICS GMBH (HOFF)

Inventor: LUDWIG W SCHLEIFER K KESSLER C RUEGER R STERN A

Language: EN

Application: CA 2052668 A 19911002 (Local application)

1

Priority: DE 4032024 A 19901009 DE 4038804 A 19901205

Original IPC: C12Q-1/68(A)
Current IPC: C12Q-1/68(A)

Germany

Publication Number: DE 4038804 A (Update 199217 E)

Publication Date: 19920416

Verfahren zur genus- oder/und spezies-spezifischen Detektion von Bakterien in einer Probenfluessigkeit

Assignee: Boehringer Mannheim GmbH, 6800 Mannheim, DE (BOEF)

Inventor: Ludwig, Wolfgang, Dr.rer.nat., 8179 Sachsenkamm, DE Schleifer, Karl-Heinz, Prof. Dr., 8044

Unterschleissheim, DE Kessler, Christoph, Dr. rer. nat., 8021 Dorfen, DE Rueger, Ruediger, Dr. med., 8124 Seeshaupt,

DE Stern, Anne, Dr. rer. nat., 8122 Penzberg, DE

Language: DE (5 pages)

Application: DE 4032024 A 19901009 DE 4038804 A 19901205 (Local application)

Original IPC: C12Q-1/68(B) G01N-33/68(B) Current IPC: C12Q-1/68(B) G01N-33/68(B)

Claim: * 1. Verfahren zur genus- oder/und spezies-spezifischen Detektion von Bak terien in einer Probenflussigkeit, **dadurch gekennzeichnet,** dass man bakterielle RNA mit einem Primer, der zu einer genus- bzw. spezies-spezifischen Region von RNA bestimmter Bakterien oder einer hochkonservie rten Region der RNA von Bakterien allgemein komplementar ist, an seinem 3prime-Ende jedoch nicht komplementar ist zur RNA von Bakterien andere r Gattung bzw. Spezies, hybridisiert, Elongation des Primers in Anwesen heit einer geeigneten Polymerase und den vier Desoxyribonukleotiden, ge gebenenfalls mit gleichzeitiger oder anschliessender Markierung des Elongationsproduktes, bewirkt und ein gebildetes Elongationsprodukt nach D enaturierung mit einem genus- bzw. spezies-spezifischen Oligonukleotid hybridisiert und die Hybridisierung uber die Markierung des Oligonukleo tids nachweist. |DE 59108610

G (Update 199721 E)

Publication Date: 19 970417

Assignee: BOEHRINGER MANNHEIM GMBH (BOEF)

Inventor: LUDWIG W SCH LEIFER K KESSLER C RUEGER R STERN A

Language: DE

Application: DE 591086 10 A 19911001 (Local application) EP 1991116745 A 19911001 (Application)

Priority: DE 4032024 A 19901009 DE 4038804 A 19901205 Related Publication: EP 480289 A (Based on OPI patent)

Original IPC: C12Q-1/68(A) Current IPC: C12Q-1/68(A)

European Patent Office

Publication Number: EP 480289 A (Update 199216 B)

Publication Date: 19920415

Verfahren zur Genus- oder/und Spezies-spezifischen Detektion von Bakterien in einer Probenfluessigkeit Method for genus or/and species-specific detection of bacteria in a testliquid Methode de detection specifique de genre ou/et d'espece de bacteries dans un liquide a prober

Assignee: BOEHRINGER MANNHEIM GMBH, Sandhofer Strasse 116, W-6800 Mannheim 31, DE (BOEF) Inventor: Ludwig, Wolfgang, Dr. rer. nat., Alpenblickstrasse 8, W-8179 Sachsenkamm, DE Schleifer, Karl-Heinz, Prof. Dr., Schwalbenstrasse 3a, W-8044 Unterschleissheim, DE Kessler, Christoph, Dr. rer. nat., Schlossbergweg 11, W-8021 Dorfen, DE RUGER R Stern, Anne, Dr. rer. nat., Karwendelstrasse 10, W-8122 Penberg, DE

Language: DE (9 pages, 0 drawings)

Application: EP 1991116745 A 19911001 (Local application) Priority: DE 4032024 A 19901009 DE 4038804 A 19901205

Designated States: (Regional Original) AT BE CH DE DK ES FR GB GR IT LI LU NL SE

Original IPC: C12Q-1/68 Current IPC: C12Q-1/68

Original Abstract: For the genus- and/or species-specific detection of bacteria in a sample liquid, bacterial RNA is hybridised with a primer which is generally complementary to a genus- or species-specific region of RNA of certain bacteria or a highly conserved region of the RNA of bacteria but is not complementary at its 3' end to the RNA of bacteria of other genus or species, the primer is elongated in the presence of a suitable polymerase and the four deoxyribonucleotides, where appropriate with simultaneous or subsequent labelling of the elongation product, an

elongation product which has formed is after denaturation hybridised with a genus- or species-specific oligonucleotide, and the hybridisation is detected via the labelling of the oligonucleotide.

Claim: * 1. Verfahren zur Genus- oder/und Spezies-spezifischen Detektion von Bak terien in einer Probenfluessigkeit, dadurch gekennzeichnet, dass man ba kterielle RNA mit einem Primer, der zu einer genus- bzw. spezies-spezif ischen Region von RNA bestimmter Bakterien oder einer hochkonservierten Region der RNA von Bakterien allgemein komplementaer ist, an seinem 3'-Ende jedoch nicht komplementaer ist zur RNA von Bakterien anderer Gatt ung bzw. Spezies, hybridisiert, Elongation des Primers in Anwesenheit e iner geeigneten Polymerase und den vier Desoxyribonukleotiden, gegebene nfalls mit gleichzeitiger oder anschliessender Markierung des Elongationsproduktes, bewirkt und ein gebildetes Elongationsprodukt nach Denatur ierung mit einem Genus- bzw. Spezies-spezifischen Oligonukleotid hybrid isiert und die Hybridisierung ueber die Markierung des Oligonukleotids nachweist. li |EP 480289 B1 (Update 199715 E)

Publication Date: 1997 0312

Verfahren zur genus- und spezies-spezifischen Detektion von Bakt erien in einer Probenfluessigkeit Method for genus and species-specific detection of bacteria in a testliquid Methode de detection specifique de genre et d'espece de bacteries dans un liquide a prober

Assignee: BOEHRINGER MANNHEIM GMBH, 68298 Mannheim, DE (BOEF)

Inventor: Ludwig, W olfgang, Dr. rer. nat., Alpenblickstrasse 8, W-8179 Sachsenkamm, DE Sch leifer, Karl-Heinz, Prof. Dr., Schwalbenstrasse 3a, W-8044 Unterschleis sheim, DE Kessler, Christoph, Dr. rer. nat., Schlossbergweg 11, W-8021 Do rfen, DE Rueger, Ruediger, Dr. med., Tutzinger Strasse 2, W-8124 Seesha upt, DE Stern, Anne, Dr. rer. nat., Karwendelstrasse 10, W-8122 Penberg, DE

6.34

Language: DE (9 pages, 0 drawings)

Application: EP 1991116745 A 19 911001 (Local application) Priority: DE 4032024 A 19901009 DE 4038804 A 19901205

Designated States: (Regional Original) AT BE CH DE DK ES FR G B GR IT LI LU NL SE

Original IPC: C12Q-1/68(A) Current IPC: C12Q-1/68(A)

Claim: 1. Verfahren zur genus- und spezies-spezifischen Detektion v on Bakterien in einer Probenfluessigkeit, mit den Schritten - bakteri elle RNA wird mit einem Primer, der zu einer genus-spezifischen Region von RNA bestimmter Bakterien oder einer hochkonservierten Region der RN A von Bakterien allgemein komplementaer ist, an seinem 3'- Ende jedoch n icht komplementaer ist zur RNA von Bakterien anderer Gattung hybridisie rt, - Elongation des Primers in Anwesenheit einer geeigneten Polymera se und den vier Desoxyribonukleotiden, gegebenenfalls mit gleichzeitige r oder anschliessender Markierung des Elongationsproduktes, - ein geb ildetes Elongationsprodukt wird nach Denaturierung mit einem spezies-sp ezifischen Oligonukleotid hybridisiert, * - die Hybridisierung wird ueber die Markierung des Oligonukleotids n achgewiesen. 1. Method for the genus-specific and species-specifi c detection of bacteria in a sample liquid comprising the steps - bac terial RNA is hybridized with a primer which is complementary to a genu s-specific region of the RNA of particular bacteria or to a highly cons erved region of the RNA of bacteria in general, but which is not complementary at its 3' end to the RNA of bacteria of another genus, - the primer is elongated in the presence of a suitable polymerase and the fo ur deoxyribonucleotides, if desired, with a concurrent or subsequent la belling of the elongation product, - an elongation product formed is hybridized with a species specific oligonucleotide after denaturation, * - the hybridization is detected by means of the oligonucleotide labe l.

Spain

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Assignee: BOEHRINGER MANNHEIM GMBH (BOEF)

Language: ES

Application: EP 1991116745 A 19911001 (Application)
Priority: DE 4032024 A 19901009 DE 4038804 A 19901205
Related Publication: EP 480289 A (Based on OPI patent)

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Japan

Publication Number: JP.4258299 A (Update 199243 E)

Publication Date: 19920914

Assignee: BOEHRINGER MANNHEIM GMBH (BOEF)

Language: JA (6 pages)

Application: JP 1991257519 A 19911004 (Local application) Priority: DE 4032024 A 19901009 DE 4038804 A 19901205

Original IPC: C12Q-1/68(A)

Current IPC: C12Q-1/68(A)|JP 1994040839 B2 (Update 199420 E)

Publication Date: 19940601

Assignee: BOEHRINGER MANNHEIM GMBH (BOEF)

Inventor: LUDWIG W SCHLEIFER K H KESSLER C RUGER R STERN A

Language: JA (6 pages)

Application: JP 1991257519 A 19911004 (Local application) Priority: DE 4032024 A 19901009 DE 4038804 A 19901205 Related Publication: JP 04258299 A (Based on OPI patent)

Original IPC: C12Q-1/68(A) Current IPC: C12Q-1/68(A)

United States

Publication Number: US 6225094 B1 (Update 200126 E)

Publication Date: 20010501

Method for the genus-specific or/and species-specific detection of bacteria in a sample liquid.

Assignee: Roche Diagnostics GmbH, Mannheim, DE (HOFF)

Inventor: Ludwig, Wolfgang, Sachsenkam, DE Schleifer, Karl-Heinz, Unterschleissheim, DE Kessler, Christoph,

Dorfen, DE Rueger, Ruediger, Seeshaupt, DE Stern, Anne, Penzberg, DE

Agent: Arent Fox Kintner Plotkin Kahn, PLLC

Language: EN

Application: US 1991772026 A 19911008 (Continuation of application) US 199373985 A 19930608 (Local

application)

Priority: DE 4032024 A 19901009 DE 4038804 A 19901205

Original IPC: C12P-19/34(A) C12Q-1/68(B) Current IPC: C12P-19/34(A) C12Q-1/68(B)

Original US Class (main): 43591.51

Original US Class (secondary): 43591.2 43591.1 4356

Original Abstract: For the genus-specific or/and species-specific detection of bacteria in a sample liquid, bacterial RNA is hybridized with a primer which is complementary to a genus-specific or species-specific region of the RNA of particular bacteria or to a highly conserved region of the RNA of bacteria in general, but which is not complementary at its 3prime end to the RNA of bacteria of another genus or species, the primer is elongated in the presence of a suitable polymerase and the four deoxyribonucleotides, if desired, with a concurrent or subsequent labelling of the elongation product and an elongation product formed is hybridized with a genus-specific or species-specific oligonucleotide after denaturation and the hybridization is detected by means of the oligonucleotide label.

Claim: 1.A method for genus-specific and/or species-specific detection of bacter ia in a sample liquid comprising the steps of: * a) hybridizing bacterial RNA with a primer which is complementary to a genus-specific region of RNA of a particular bacteria wherein the p rimer is not complementary at its 3prime end to RNA of bacteria from another genus, * b) elongating said primer in the presence of a suitable polymerase and deoxyribonucleotides to form an elongation product, * c) hybridizing said elongation product after denaturation with a labe led oligonucleotide which is specific for a particular species within a genus of bacteria, and * d) detecting any hybridization by means of the labeled eligonucleotid e.

-iA

South Africa

Publication Number: ZA 199108024 A (Update 199235 E)

Publication Date: 19920729

Assignee: BOEHRINGER MANNHEIM GMBH (BOEF)

Inventor: KESSLER C RUEGER R SEIBL R KRUSE-MUELLER C BERNER S

Language: EN (47 pages)

Application: ZA 19918024 A 19911008 (Local application)

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